

APR 01 2004

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

RE: Application of

Joseph LOSCALZO et al.

Serial No. 10/687,706

Filed: October 20, 2003

For: METHODS OF TREATING VASCULAR DISEASES CHARACTERIZED BY NITRIC OXIDE INSUFFICIENCY

Honorable Commissioner for Patents
Alexandria, VA 22313-1450: Attn: Office of Initial
: Patent Examination

: Group Art Unit: 1651

:

REQUEST FOR CORRECTED FILING RECEIPT

Attached is a copy of the Filing Receipt received from the U.S. Patent and Trademark Office in the above-referenced application. The Filing Receipt is missing some of the priority information. Also attached is a copy of the first page of the specification listing the correct priority information. Please add the Priority information as indicated in red on the attached copy of the Filing Receipt. It is requested that a corrected filing receipt be issued.

The domestic priority data on the Official Filing Receipt currently reads:

This application is a CIP of 09/697,317 10/27/2000 PAT 6,635,273
which claims benefit of 60/179,020 01/31/2000
and claims benefit of 60/162,230 10/29/1999

Please correct the domestic priority data on the Official Filing receipt to recite:

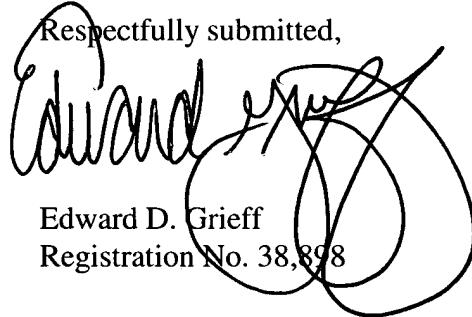
**This application is a CIP of 09/697,317 10/27/2000 PAT 6,635,273
which claims benefit of 60/179,020 01/31/2000
and claims benefit of 60/162,230 10/29/1999**

**This application is a continuation of 10/415,136 04/25/2003
which is a 371 of PCT/US01/14245 05/02/2001
which claims benefit of PCT/US00/29528 10/27/2000**

The present application properly claims priority to 6 different applications (2 US nonprovisional applications, 2 PCT applications and 2 provisional US applications). Applicants respectfully request that the Official Filing Receipt be appropriately corrected.

Request for Correct Filing Receipt
US Application No. 10/687,706
Page 2 of 2

Since the error was made by the U.S. Patent and Trademark Office, it is believed that no fee is required with regard to this Request; however, in the event that a fee is required, the Commissioner is authorized to charge such fee to Deposit Account No. 08-0219. A duplicate copy of this letter is attached.

Respectfully submitted,

Edward D. Grieff
Registration No. 38,898

Date: March 31, 2004

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**UNITED STATES PATENT AND TRADEMARK OFFICE**

Page 1 of 2
magn 3/17
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APPL NO.	FILING OR 371 (c) DATE	ART UNIT	FIL FEE REC'D	ATTY.DOCKET NO	DRAWINGS	TOT CLMS	IND CLMS
10/687,706	10/20/2003	1651	2160 O P F JC174	102258.170 US2	3	169	7

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EDWARD D GRIEFF
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**CONFIRMATION NO. 2830****CORRECTED FILING RECEIPT**

OC000000012096914

Date Mailed: 03/12/2004

Receipt is acknowledged of this regular Patent Application. It will be considered in its order and you will be notified as to the results of the examination. Be sure to provide the U.S. APPLICATION NUMBER, FILING DATE, NAME OF APPLICANT, and TITLE OF INVENTION when inquiring about this application. Fees transmitted by check or draft are subject to collection. Please verify the accuracy of the data presented on this receipt. If an error is noted on this Filing Receipt, please write to the Office of Initial Patent Examination's Filing Receipt Corrections, facsimile number 703-746-9195. Please provide a copy of this Filing Receipt with the changes noted thereon. If you received a "Notice to File Missing Parts" for this application, please submit any corrections to this Filing Receipt with your reply to the Notice. When the USPTO processes the reply to the Notice, the USPTO will generate another Filing Receipt incorporating the requested corrections (if appropriate).

Applicant(s)

Joseph Loscalzo, Dover, MA;
Joseph A. Vita, Hingham, MA;
Michael D. Loberg, Boston, MA;
Manuel Worcel, Boston, MA;

Domestic Priority data as claimed by applicant

This application is a CIP of 09/697,317 10/27/2000 PAT 6,635,273
which claims benefit of 60/179,020 01/31/2000
and claims benefit of 60/162,230 10/29/1999

*This application is a continuation of 10/415,136, 4/25/2003
which is a Section 371 national phase of PCT/US01/14245, 5/2/2001
Foreign Applications which claims benefit of PCT/US00/29528, 10/27/2000*

If Required, Foreign Filing License Granted: 01/20/2004

Projected Publication Date: 04/29/2004

Non-Publication Request: No

Early Publication Request: No

** SMALL ENTITY **

HALE AND DORR LLP Wash, DC

Proj. Date: 1/22/08. 170 US2

Proj. Due Date: _____

Proj. Ctr. No.: _____

Final Date: _____

Ext. _____ Dk'd By VC

Title

Methods of treating vascular diseases characterized by nitric oxide insufficiency

Preliminary Class

424

**LICENSE FOR FOREIGN FILING UNDER
Title 35, United States Code, Section 184
Title 37, Code of Federal Regulations, 5.11 & 5.15**

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Methods of Treating Vascular Diseases Characterized by Nitric Oxide Insufficiency

Related Applications

This application is (i) a continuation-in-part of U.S. Application No. 09/697,317 filed October 27, 2000, which claims priority to U.S. Application No. 60/179,020 filed January 31, 5 U.S. Application No. 60/162,230 filed October 29, 1999; and (ii) a continuation of U.S. Application No. 10/415,136 filed April 25, 2003, which is a § 371 of PCT/US01/14245 filed May 2, 2001, which claims priority to PCT/US00/29528 filed October 27, 2000.

Field of the Invention

The present invention provides methods of treating and/or preventing vascular diseases 10 characterized by nitric oxide insufficiency by administering a therapeutically effective amount of at least one antioxidant or a pharmaceutically acceptable salt thereof, and at least one compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase, and, optionally, at least one nitrosated angiotensinconverting enzyme inhibitor, 15 nitrosated beta-adrenergic blocker, nitrosated calcium channel blocker, nitrosated endothelin antagonist, nitrosated angiotensin II receptor antagonist, nitrosated renin inhibitor, and/or at least one compound used to treat cardiovascular diseases. The antioxidant may preferably be a hydralazine compound or a pharmaceutically acceptable salt thereof. The compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived 20 relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase may preferably be isosorbide dinitrate and/or isosorbide mononitrate. The present invention also provides methods of treating and/or preventing vascular diseases characterized by nitric oxide insufficiency by administering a therapeutically effective amount of at least one nitrosated angiotensinconverting enzyme inhibitor, nitrosated beta-adrenergic blocker, nitrosated 25 calcium channel blocker, nitrosated endothelin antagonist, nitrosated angiotensin II receptor antagonist and/or nitrosated renin inhibitor, and, optionally, at least one antioxidant and/or at least one compound used to treat cardiovascular diseases. The present invention also provides methods of treating and/or preventing Raynaud's syndrome by administering a therapeutically effective amount of at least one antioxidant or a pharmaceutically acceptable salt thereof, and at 30 least one compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase, and, optionally, at least one nitrosated angiotensin-